## SPECIAL ARTICLE

# Cancer pain... who cares? International and national patterns of evidence-based global guidelines recommendations for physicians on the Web (2011 vs. 2018)

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### Summary

**Purpose:** Although pain is a common event during treatment of cancer, its assessment and management remains suboptimal in everyday clinical practice at global level.

Methods: Considering both the important role of internet in daily life and that clinical guidelines are important for translating evidence in clinical practice, we performed a prospective study to scrutinize the magnitude of updated evidence-based cancer-pain guideline recommendation for physicians on the web. Changes over-time at a global level were scrutinized at two time points: 2011 for baseline and 2018 at first follow-up. Both anesthesiology and oncology societies were analyzed.

Results: In 2011 we scrutinized 181,00 WebPages and 370 eligible societies were identified: 364 of these were eligible for analyses both in 2011 and 2018. The magnitude of cancer pain updated and evidence-based guideline recommendations on the web for health care providers was extremely low

at global level and at any time point considered: 1.1% (4/364) in 2011 and 4.7% (17/364) in 2018. Continental and intercontinental patterns, National's highest developmental index, oncology tradition and economic-geographic areas were not found to influence cancer pain web-guideline provision. In 2018, pain & supportive care societies provided the highest rate of updated evidence-based cancer-pain guidelines for clinicians. Only 3/25 medical oncology societies and 1/34 radiation oncology societies, provided own or e-link (to other societies') evidence-based guidelines in their websites.

**Conclusions:** Major medical oncology and radiation oncology societies - at global level - fail to produce updated cancer pain recommendations for their physicians, with most of these providing no or inconsistent or outdated guidelines.

Key words: cancer pain, global awareness, guideline imple*mentation, web, medical societies, oncology, anesthesiology* 

# Introduction

cacy have been cumulatively achieved across decades and major survival improvements in both palliative strong determinant of rationalized oncology prac-

Stepwise improvements in cancer treatment effi- and radical treatment settings have been reached. Nonetheless, cost-effectiveness remains a

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tice. Billions are spent each year for prematurely approved costly treatments of uncertain benefit [1] as well as and for genomic diagnostic tests of equivocal utility [2], with most of these being potentially beneficial only for very few and selected group of patients.

But what happens with cancer pain? It affects more than half of cancer patients, with a prevalence of 55% among patients on anticancer treatment, 66.4% among those with advanced metastatic or terminal phases of the disease, and 39.3% following curative treatment [3]. Moderate to severe pain is reported by 38% of all patients [3], with severe impact on quality of life and performance of normal daily activities [4,5]. To date, one third of the patients still do not receive pain medication proportional to their pain intensity levels [6]. Pain research, pain assessment and management remain suboptimal in everyday clinical practice with half of the patients believing that their quality of life is not considered a priority in their overall care by their health care professionals [5,7]. Thus, cancer pain is a serious public health problem and a major concern for more than 10 million people yearly diagnosed with cancer worldwide [8].

How to ameliorate the management of cancer pain, and how to improve and assist the physicians' awareness in cancer pain management in daily clinical activities remains a hostile cornerstone to be solved.

Clinical practice guidelines are important for translating evidence in medical decision making and reducing undesirable practices encouraging services of proven efficacy [9]. Medical guidelines/ recommendations provision in websites has been of extreme importance in improving patients' safety, reducing complications and shortening the length of stay among Medicare beneficiaries [10].

Since most recognized medical societies have very extensive membership, organize a large number of educational meetings worldwide, and have substantial influence upon their members, subscribers, and visitors; we hypothesized that one of the possible causes of current medical mismanagement of cancer pain might stem from a low number of web guidelines implementation among oncology, educational and policymaker medical societies. Thus, we set to examine the global coverage of cancer pain guidelines recommendations on the web for clinicians produced by international and national oncology societies. The magnitude of cancer pain guideline production on the web and its changes over time (2011 vs 2018 estimates) were scrutinized in a prospective web-based study. Since different level of development and economy

might largely influence clinical daily practice and priorities in guideline implementation, we further separately scrutinized differences in cancer pain guideline implementation among the 10 highest developed countries [11], the 10 countries with long lasting tradition in medical oncology, and 6 different economic-geographic areas.

#### Methods

#### Identification of pertinent societies and caregivers

In 2011, 181,200 WebPages were scrutinized in order to identify anesthesiology, oncology and pain societies/organizations that might have provided web guidelines regarding cancer pain. We retrieved both international societies (intercontinental, African, Asian, European, Oceanian, North American, South American) and national organizations belonging either to one of the top 10 countries with the highest development index (Norway, Australia, New Zealand, USA, Ireland, Liechtenstein, Netherlands, Canada, Sweden, Germany) [11], or to 10 countries with a long lasting tradition in medical oncology but not included in the top 10 high developed countries (Austria, Belgium, China, Denmark, France, Japan, Italy, UK, Spain, Switzerland) [appendix\_1 methods]. Due to notable economy and development differences between South and North American countries, the continental entities were separately searched and analyzed for North and South America.

National associations identified were further grouped by geographical-economic areas: Australia-New Zealand, Benelux, United Kingdom of Great Britain and Ireland, German speaking countries, North American, Scandinavian, South European and East Asian countries. Further methodological details are reported in Appendix\_1 methods (Table 1).

Web searches identified 370 potentially eligible societies. Since one society was double reported and 5 societies ceased, 364 societies/organizations were eligible for analyses. (Figure 1. research flow chart). (appendix 2. List of analysed societies).

Screening of the 364 eligible societies' web-sites for guideline recommendations was performed in June 2011 and in June 2018.

#### Outcomes

To scrutinize the global magnitude of "updated" and "evidence-based" guideline recommendations for cancer pain for physicians on the web and its changes over time. We considered as "updated" all the web guidelines that have been produced or revised or lastly adjourned within the last 5 years. Evidence-based were considered all guidelines including randomized controlled trials and/ or meta-analyses in their references. Furthermore, we considered eligible only cancer pain guideline pertaining the general assessment and management of cancer pain. "Solo" specialist guideline (such as "solo" radiation protocols for bone pain among radiotherapy societies, or "solo" intrathecal use of opioids among anesthesiology societies) were not included in the final analyses.



Figure 1. Research flow-chart.

### Results

A statistical improvement in the production of evidence-based and updated web-recommendations for cancer pain was observed over time (2011 vs 2018) at global level (x<sup>2</sup>=8.2866, p=0.039). Nonetheless, the magnitude of recommendation provision was inquiringly low for any outcome considered. Only 16 and 44 societies / health providers provided some form of cancer pain web-recommendations (any setting considered) in 2011 and 2018; of these, only 4 societies in 2011 [12-15] and only 17 in 2018 [16-32] were providing "updated and evidence-based" cancer pain recommendations for physicians in their web sites (Table 1). Thus, the proportion of medical societies implementing cancer pain updated evidence-based guidelines for clinicians were almost null either in 2011 and 2018 (1.1 vs 4.7%).

In 2011, the 4 societies provided recommendations both for cancer pain assessment and cancer pain treatment [12-15], while in 2018 all the 17 societies were providing recommendations for cancer pain treatment [16-32], but only 10 provided recommendations for cancer pain assessment [21,23, 24-27,30-32].

At their best (2018 analyses), only one intercontinental [16], 3 European [17,18,26], 4 North American [19-22], 3 Dutch [27-29], 2 German [30,31], 1 Japanese [32], 1 Italian [24], 1 Spanish [25], 1 UK [23], and 3 US medical societies (3/56) [20-22] were providing evidence-based updated guidelines for cancer pain in their web sites. No evidence-based updated recommendations for clinicians were found across African, Asian, Oceanian, South American medical societies, and the societies analyzed of the resting 13 countries (Table 1). Guideline release for clinicians was not influenced by the continent analyzed, the national high developmental index and the national high oncology tradition. Similarly, when the countries were grouped and analyzed by economic-geographic areas (Australia-New Zealand vs Benelux vs German speaking countries vs North American vs Scandinavian vs South European vs Great Britain and Ireland vs East Asian), no statistical differences were found in the proportion of societies providing updated evidence-based web-recommendation for cancer pain (2011: Yates' x<sup>2</sup>=3.719, p=0.811; 2018: Yates' x<sup>2</sup>=3.429, p=0.843).

Only the society type (anesthesiology vs oncology vs supportive care & pain societies) was found to influence cancer pain web-guideline provision in 2018 (Yates'  $x^2$ =6.994, p=0.030). Nonetheless, despite a higher proportion of evidence-based updated web cancer pain guidelines was found among pain and supportive care medical societies (16.6%, 4/24) [16-19], while web-guideline delivery among oncology and anesthesiology societies did not overcome the 4.3% (Table 1).

When the societies' sub-types were analyzed, a higher proportion of web recommendations for physicians was evident for supportive care (30%) [17-19], and medical oncology societies (12%) [21,24,25], and were null or almost null across other societies subtypes (Table 1).

	Eligible n=364	2011 any Recomm. Cancer pain n=16	2018 any Recomm. Cancer pain n=44	2011 EB.U Guidelines Cancer pain n=4	2018 EB.U Guidelines Cancer pain n=17
Continent					
Intercontinental	54	2	5	1	1
North America	69#	6#	12#	1#	4#
South America	6	1	1	0	0
Europe	35	1	5	1	3
Africa	11	1	1	0	0
Asia	5	0	0	0	0
Oceania	2	0	0	0	0
Top 10 Developed Countries*					
Norway	4	0	0	0	0
Australia	16	0	1	0	0
New Zealand	7	0	0	0	0
USA	52	5	11	1	4
Ireland	10	0	0	0	0
Liechtenstein	0	0	0	0	0
Netherlands	9	0	3	0	3
Canada	17	1	1	0	0
Sweden	4	0	0	0	0
Germany	10	0	2	0	2
Other countries					
Japan	13	0	2	0	1
United Kingdom	18	3	6	1	1
Italy	11	2	4	0	1
Switzerland	14	0	0	0	0
Spain	13	0	1	0	1
Belgium	9	0	1	0	0
Denmark	7	0	0	0	0
France	12	0	0	0	0
China	15	0	0	0	0
Austria	10	0	0	0	0
Geographic economic area	27	0	1	0	0
Australia - New Zeal.	23	0	1	0	0
Benelux	18	0	4	0	3
Germanophone North American	34	0	2	0	2
Scandinavian	69 15	6	12	1	4
	15 36	0	0	0	0 2
South European UK-Ireland	28	2 3	5	0	
East Asian	28 28	0	6 2	1 0	1
Society, category	20	U	2	U	Ţ
Anesthesia	79	2	6	0	1
Oncology	256	2	25	3	1
Pain & Supportive Care	250 24	4	11	1	4
Other	24 5	4	2	0	4
Continued on the next nage	0	1	2	U	1

### Table 1. Demographics of the scrutinized societies and caregivers organizations

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	Eligible n=364	2011 any Recomm. Cancer pain n=16	2018 any Recomm. Cancer pain n=44	2011 EB.U Guidelines Cancer pain n=4	2018 EB.U Guidelines Cancer pain n=17
Society, subtype					
Anesth. Comprehen.	45	1	5	0	1
Anesth. Other	34	1	1	0	0
Pain	14	3	6	1	1
Cancer Research	52	1	2	0	0
Radiation Oncology	34	0	5	0	1
Medical Oncology	25	2	5	1	3
Surgical Oncology	15	0	1	0	1
Supportive Care	10	1	5	0	3
Compr. CA. MGM**	71	5	10	2	5
Other Societies	64	2	4	0	2

Distribution of the scrutinized societies and caregivers organizations by location, type (anesthesiology, oncology, pain); eligibility, accessibility and relative guideline recommendations.\* Countries were selected from the top 10 countries from the human development index; \*\*Compr. Cancer Management, "North American guidelines were obtained by the addition of USA+ Canada societies/organizations.

### Discussion

Our analysis provides strong evidence for lack of web-guidelines for physicians for the assessment and management of cancer-related pain. Astonishingly, even in 2018, in our study, only 6.8% (4/59) of the "gate-keepers" specialties (medical oncology and radiation oncology) involved in the management of cancer patients provided evidencebased updated recommendation for cancer pain in their web-sites. Of note, ASCO (American Society of Clinical Oncology - the major medical oncology society worldwide) provided evidence-based pain guidelines only for the restricted sub-setting of cancer survivors [21], while no guidelines were provided for patients with active disease in any setting considered (under treatment, under follow-up or palliation). At the same time, ESMO (European Society of Medical Oncology - the European counterpart for medical oncology and second worldwide provider) presented outdated guidelines, thought new cancer pain guidelines were in press at the time of writing of our report [33].

Neither ASTRO (American Society for Therapeutic Radiology and Oncology), nor ESTRO (European Society for Therapeutic Radiology and Oncology), provided relative recommendations in their web sites (Table 1).

Cancer pain is a major public health problem. The crucial question is why this low level of priority exists, especially when the prevalence of cancer-related pain appears to be very high, and considering that it may severely jeopardize quality of life and performance of normal daily activities [3-5]. In some cases, patients may fear pain more than potential death from their cancer and this fear has aided the drive for the agenda of physiciansassisted suicide [34].

For all the above-mentioned threats the World Health Organization (WHO) developed guidelines to assist in the management of cancer pain more than 30 years ago [35]. Nonetheless, the cancer pain threat is far from being solved. To date, one third of the patients still did not receive pain medication proportional to their pain intensity levels [6], pain research, pain assessment and management remain suboptimal in everyday clinical practice with half of the patients believing that their quality of life is not considered a priority in their overall care by their health care professional [5-7]. Consequently, cancer patients' dissatisfaction is very high. Inevitably, patients and their family members are prone to find their solutions by themselves, frequently by surfing blindly in the internet [36]. Nonetheless, these blind internet searches are of particular threat and may jeopardize the same patients' outcomes since the cancer pain and cancer-cachexia Web information is largely dominated by the extremely strong market of para-medicine and counterfeit drugs [36,37].

Recently, the European Association for Palliative Care (EAPC) defined the untreated cancer pain as "scandal of global proportion", as a combined action of EAPC, European Society of Medical Oncology (ESMO), the Pain Policy Studies Group (PPGS), the Union International Cancer Control (UICC) and the WHO underscored a lack of access to opioids medication at global level [38].

Can scarcity in guideline implementation modify medical thought in decision-making and generate deficits in cancer pain assessment / management in daily practice? to establish guidelines on a certain subject?

An impressive number of medical, anesthesiology, and oncology societies have been developed over time and are engaged in providing flourishing professional and scientific activities. Many of these organizations have extensive membership bases and organize large meetings. In 2018, half of these societies provided guidelines, recommendations and position statements within their websites that have substantial influence upon their members, subscribers, and websites visitors [39-41]. Web clinical practice guidelines are important for translating evidence in medical decision-making and clinical practice applications, reducing undesirable practices, encouraging services of proven efficacy, improving patients' safety and reducing complications [10]. Nonetheless, billions of dollars are spent each year for guidelines recommending prematurely approved costly treatment of uncertain benefit [1] and for guideline recommending genomic diagnostic tests of equivocal utility [2]. One may thus wonder why these societies do not prioritize guideline implementation for a pivotal and common problem (cancer pain) in daily clinical practice?

Diverging causes, such as scarce funding, lack of motivation, lack of impact on professional development, no interest from stakeholders, conflicting roles or educational deficits in pre- or postgraduate settings should be examined. Thus, in some clinical situations, the flourishing of high professional activity of medical societies might not be translated in an equal benefit for patients.

Prevalence and severity of a determinate clinical entity, as well as the patients' expectancies from physicians might be substantially different from priorities of physicians and medical professional societies. Nowadays oncologists resemble more and more to molecular biologists, while patients are more and more seeking for a doctor who cares for them.

How to solve these discrepancies? To positively impact the development of clinical practice guidelines and put them on the web might represent a new challenging field for the future.

Our study presents some limitations. First of all, since there are no established validated searches for unearthing professional societies and organizations, some of them may have been missed by our searches. However, given the multiple layers of our search, and the large number of oncology societies retrieved, it is unlikely that prominent entities were missed and that missed societies

What triggers a determined medical society might change the global patterns of web-guideline provision. Details on this study methodology had been already published in the literature [36,37,42-44]. Secondly, the human development index (HDI) changes over time. Thus, in June 2018 (at the time of data extraction) [45], countries' position varied compared to the top 10 positions available in June 2011 [11]. Among the 188 nations analyzed by the HDI, 7 countries included of the top 10 HDI at the time of our analyses in the 2011 (Norway, Australia, USA, Ireland, Netherland, Canada, Germany) [11] continued to be in the top 10 at the time of our data extraction in June 2018 [45]. The remaining three countries continue to rank at the top of the list, all included in the top 15 positions (New Zealand 13/188, Sweden 14/188, and Liechtenstein 15/188) [45]. Thereafter, no significant biases may be attributed to country highest developmental national index migration at the two time-point of analyses.

> In conclusion, our study outlined that overall cancer pain updated and evidence-based webguidelines for physicians are remarkably scarce despite they seem to improve overtime. Moreover, the phenomenon is independent of continent, developmental index of the nation analyzed, oncology tradition and economic-geographic area.

> Despite the fact one third of cancer patients did not receive adequate pain medications, medical oncology and radiotherapy societies fail to regularly produce updated cancer pain recommendations for their physicians, with most of these providing no or inconsistent or outdated guidelines. Cancer pain, who cares?

#### **Conflict of interests**

Study concept and study design (DM, NPP, AV, KK); internet screening for available societies in 2011 (TL, KK); scrutiny of relative web sites in 2011 (TL, KK); discussion upon scrutiny uncertainties 2011 (DM, NPP, AV, TV); protocol review and amendment in 2017 (DM, NPP, AV, GZV, TL, KK, TV, PN); scrutiny of relevant web sites in 2018 (PF, CG, GZF, MY); discussion upon scrutiny uncertainties in 2018 (NP, FK); critical appraisal of the results 2011, 2018 and analyses (DM, GP, GZV, DV, EP, ET, PN); manuscript writing (DM, GZV); Final details in manuscript editing: GP.

#### **Conflict of interests**

Authors have no conflicts of interest and no financial interest to declare.

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### Appendix 1

#### Methods

Identification of pertinent societies and caregivers

In 2011 we constructed a database of anesthesiology, oncology and pain societies/organizations (educational, professional, health policymaker, caregivers) that might provide guidelines for cancer pain. We considered societies and organizations that were intercontinental (with a global outlook), continental (including two or more countries in the same continent), or national belonging to one of the top 10 countries with the highest development index [1].

Countries with a long lasting tradition in medical oncology (countries in which were performed the largest number of chemo/hormonal therapy randomized trials for advanced malignancies, based our previous meta-analyses [2-5]) but not included in the top 10 high developed countries, were further included in the internet searches (Table of main manuscript).

We performed internet searches (last search June 2011) involving possible combinations of 11 subject matters ("anesthesiology", "anesthesiological", "cancer", "oncology", "medical oncology", "clinical oncology", "radiation oncol-ogy", "radiotherapy", "surgical oncology", "cancer research", "supportive oncology"), 3 terms for educational and policymaker societies ("society" or "association" or "organization") and 30 terms of geographic identifiers (10 pertaining to continents: "Asian", "American", "North American", "South American", "America Latina", "African", "European", "Australian", "Oceania", "International"; 10 pertaining to eligible countries by the highest development index [1]: "Norway", "Australia", "New Zealand", "USA", "Ireland", "Liechtenstein", "Netherlands", "Canada", "Sweden", "Germany"; and 10 pertaining to countries with a long lasting tradition in oncology but not included in the top 10 high developed countries: "Austria", "Belgium", "China", "Denmark", "France", "Japan", "Italy", "UK", "Spain", "Switzerland "). Due to notable economy and development differences between South and North American countries, the continental entities were separately searched and analyzed for North and South America [1]. This methodology for the identification of pertinent societies and caregivers had been previously used and described [6-8].

National associations identified were further grouped by geographical-economic areas a) Australia-New Zealand, b) Benelux (Belgium and Netherland), c) Germanophone (Austria, Germany, Liechtenstein Switzerland), d) North American ( US and Canada), e) Scandinavian (Denmark, Norway and Sweeden), f) South European (France, Italy and Spain), g) United Kingdom of Great Britain and Ireland, h) East Asian (Japan and China).

The first 100 results for each internet search were scrutinized. We included both societies with accessible web pages, as well as those whose presence was mentioned in some URL but did not have a webpage or their link was not functional (under construction or not working).

#### Outcomes

To scrutinize the global magnitude of updated and evidence-based guideline recommendations for cancer-pain for physicians on the web and its changes over-time. Both anesthesiology and oncology societies were analyzed.

We considered as "updated" all the web guidelines that have been produced within five years or the web page should have been reviewed or lastly adjourned within five years by the implementing organization. If time period was higher than five years we considered the guidelines as outdated. Evidence-based were considered all guidelines including randomized controlled trials and/or meta-analyses in their references.

Furthermore, we considered eligible only cancer pain guideline pertaining the general assessment and management of cancer pain. "Solo" specialist guideline (such as "solo" radiation protocols for bone pain among radiotherapy societies, or "solo" intrathecal use of opioids among anesthesiology societies) were not included in primary outcome analyses.

#### Data extraction from eligible website

From each pertinent anesthesiology / oncology / pain society and caregiver website we recorded its name, the URL, continent and/or country, sub-specialty setting (anesthesia research, comprehensive anesthesia managing, pain, supportive oncology, medical oncology, surgical oncology, radiation oncology, cancer research) and whether they provided any guideline on any subject matter (any setting) and on cancer pain related guideline (last update for baseline screening june 2011, last updated for first interim analyses june 2018, next analyses are programmed for 2025). Whenever there was availability to perform electronic battle-searches within the website, we used the terms "guidelines» or "recommendations" or "position statements" in English. For non English websites, we translated these terms into the language the website used.

Whenever any eligible guidelines were available, we recorded whether recommendations were freely accessible through the website and whether they provided separate information developed by the society/organization itself or a link to another society/organization's guidelines.

For each cancer pain guideline retrieved, we further addressed if it was implemented for patient or for physicians, whether it pertained cancer pain assessment or treatment setting. In order to evaluate guidelines consistency we further extract whether references were provided to support the guidelines statements, whether the evidence from randomized controlled trials and/or meta-analyses were provided to support the guidelines statements.

At each time point of analyses (2011, 2018) at first screening, we did not use a strict definition for guideline and any kind of recommendation ("guidelines" or "recommendations" or "position statements" or "suggestions" or "indications") both for patients and physicians were recorded. However in the analyses for primary outcomes only evidenced

#### References

- 1. Human development reports. Human development index 5. (HDI) Rankings.
- Mauri D, Polyzos NP, Salanti G, Pavlidis NP, Ioannidis JP. Multiple treatment meta-analysis of chemotherapy and targeted treatment therapies in advanced breast cancer. J Natl Cancer Inst 2008;100:1780-91.
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based and updated guidelines for physicians were considered of value. Guidelines from web pages to be updated for more than five years were considered outdated.

Since all medical societies may have not the possibility to produce "own" guidelines, and considering that medical societies in their websites may provide guidelines either as "own produced guidelines", either as a "link" to guidelines produced by other medical societies, we considered of value both guideline produced by "own" and/or as a "link" to a specific web site of another society with web recommendation for cancer pain.

#### Analyses

We evaluated whether the proportion of associations/ organizations present intercontinental and international variations and the possible role played by the society type and subtype in guideline implementation. Group comparisons for categorical variables used chi-square, Fisher's exact test, and Yates'  $x^2$ . Whenever data scarcity was too high to allow analyses we used descriptive statistics.

#### Protocol amendment in 2017

In November 2017, considering the scarcity of updated cancer pain web-guidelines available, the board of primary investigators decided to recognize as updated all guidelines produced or adjourned within a period of five years (in the initial protocol updated guidelines/recommendations should have been provided or adjourned within a period of three years).

- Mauri D, Pavlidis N, Polyzos NP, Ioannidis JP.Survival with aromatase inhibitors and inactivators versus standard hormonal therapy in advanced breast cancer: metaanalysis. J Natl Cancer Inst 2006;98:1285-91.
- Polyzos NP, Valachis A, Mauri D, Ioannidis JPA. Industry involvement and baseline assumptions of cost-effectiveness analyses: diagnostic accuracy of the Papanicolaou test. CMAJ 2011;183:E337-43.
- Polyzos NP, Mauri D, Ioannidis JPA. Guidelines on chemotherapy in advanced stage gynecological malignancies. An evaluation of 224 professional societies and organizations. PLoS One 2011;6:e20106.
- 8. Mauri D, Tsiara A, Valachis A et al. Cancer Cachexia: Global awareness and guidelines implementation on the web. BMJ Support Palliat Care 2013;3:155-60.

### Appendix 2

#### List of the 364 societies/organizations scrutinized Age Anaesthesia Association

World Federation Societies of Anesthesiologists ACORN CRO Africa Oxford Cancer Consortium African Cancer Organization African Organisation for Research and Training in Cancer African Radiation Oncology Group African Women's Cancer Awareness Association Age Anaesthesia Association Alles Over Cemotherapie Alliance mondiale contre le cancer American Academy of Pain Management American Anti-Cancer Society American Association for Cancer Education American Association for Cancer Research American Brachytherapy Society American Cancer Society

American College of Oncology Administrators
American College of Radiation Oncology
American Institute for Cancer Research
American Pain Society
American Society for Therapeutic Radiology and Oncology
American Society of Anesthesiologists
American Society of Clinical Oncology
American Society of Preventive Oncology
American Society of Regional Anesthesia and Pain Medicine
American-Italian Cancer Foundation
Anaesthesia Patient Safety Foundation
Anaesthetic Research Society
Arbeitsgemeinschaft Internistische Onkologie
Asian American Network for Cancer Awareness
Asian Clinical Oncology Society
Asian Federation of Organizations for Cancer Research and
Control
Asian Fund for Cancer Research
Asian- Oceanian Clinical Oncological Society
Asian Pacific Organization of Cancer Prevention
Association for Directors of Radiation Oncology Programs
Association for International Cancer Research
Association for Research on Treatment against Cancer
Association for the International Development of Anesthesia
Association Latin American for Therapeutic Radiation Oncol-
ogy (ALATRO)
Association of Physician Assistants in Oncology
Association of American Cancer Institutes
Association of Anesthesia Clinical Directors
Association of Burns and Reconstructive Anaestheists
Association of Cancer Executives
Association of Community Cancer Centers
Association of European Cancer Leagues
Association of Freestanding Radiation Oncology Centers
Association of Integrative Oncology and Chinese Medicine
Association of Residents in Radiation Oncology
Association of University Anesthesiologists
Associazione Anestesisti Rianimatori Ospedalieri Italiani
Australasian Society of Anaesthesia Paramedical Officers
Australian Cancer Research Foundation
Australian Society of Anaesthetists
Austrian Cancer Aid Society
Austrian cancer association
Austrian Society of Anaesthesiology, Resuscitation and In-
tensive Care
Austrian Society of Hematology and Oncology
Austrian Society of Oncology
Austrian Society of Oncology Pharmacy
Austrian Society of Radiation Oncology
Austrian Society of Natiation Oncology Austrian Society of Surgical Oncology
Belgian Association for Cancer Research
Belgian Association for Radiotherapy and Oncology
Belgian Federation Against Cancer
0 0
Belgian Pain Society
Belgian Society of Medical Oncology
Belgian Society of Surgical Oncology
Berufsverband Deutscher Anaesthesisten
British Accelerator Science and Radiation Oncology
Consortium
British Anaesthetic & Recovery Nurses Association
British Association of Cancer Research
British Association of Cancer United Patients
British Association of Surgical Oncology
British Association of Surgical Oncology British Oncological Association British Oncology Pharmacy Association

Canadian Association of General Practitioners in Oncology Canadian Association of Medical Oncologists Canadian Association of Nurses in Oncology Canadian Association of Pharmacy in Oncology Canadian Association of Provincial Cancer Agencies Canadian Association of Radiation Oncologists Canadian Cancer Action Network Canadian Cancer Advocacy Network Canadian Cancer Research Alliance Canadian Cancer Society / National Cancer Institute of Canada Canadian Oncology Societies Canadian Partnership Against Cancer Canadian Society for Surgical Oncology Cancer Advocacy Coalition of Canada Cancer assistance network Cancer Association of South Africa Cancer Australia Cancer care,Inc. Cancer Control New Zealand Cancer Council Australia Cancer Cure Foundation Cancer Federation Inc. Cancer Foundation of China / FORMER= Chinese Cancer Research Foundation Cancer Hope Network **Cancer Patients Foundation** Cancer Project Cancer research foundation of America Cancer Research Initiative of South Africa Cancer Research Institute Cancer Research Society of Canada Cancer Research UK Cancer Society of New Zealand Cancer Support Association of Western Australia Cancer Support France Cancer Trials New Zealand Cancérologues Sans Frontières" / "Oncologists Without Borders Canteen Ireland Central European Cooperation Oncology Group China East Radiation Oncology Group Chinese American Society of Anesthesiology Chinese Anti-Cancer Association Chinese cancer research foundation (China) Chinese Center for Disease Control and Prevention Chinese Medical Association Chinese Medical Association Society of Oncology Chinese Oncology Society (Taiwan) Chinese Preventive Medicine Association Chinese Society of Anesthesiologists Chinese Society of Clinical Oncology Chinese Society of Therapeutic Radiology and Oncology / Chinese Society of Radiation Oncology Clinical Cancer Research Center Clinical Oncology Society of Australia Coc Member Organization Cancer Care Initiatives Community oncology alliance Complementary and Alternative Medicine for Cancer Confederación Latinoamericana de Sociedades de Anestesiología Confederation of European National Societies of Anaesthesiologists Conseils pour la chimiothérapie Cris Foundation for Cancer Research

Cure Cancer Australia Foundation Danish Anaesthesiological Organisation Danish Cancer Society Danish Research School in Molecular Cancer Research Danish Society of Intensive Care Therapy Danish Society of Anaesthesiology and Intensive Care Medicine Danish Society of Medical Oncology Dansk Selskab for Cancerforskning Researc Deutsche Gesellschaft für Anästhesiologie und Intensivmedizin Deutsche Interdisziplinäre Vereinigung für Intensiv- und Notfallmedizin Dutch Association of Medical Oncology Dutch Association of Oncology Nurses Dutch Belgian Hemato-Oncology Cooperative Group Dutch Cancer Society Dutch Society for Radiotherapy and Oncology Dutch Society of Oncology Dutch Society of Surgical Oncology Eastern Cooperative Oncology Group European (Spain) Website of Anaesthesia, Intensive Care and Pain Medicine European Academy of Anaesthesiology European Association for Cancer Education European Association for Cancer Research European Cancer Organisation European cancer prevention organization European Masters Program in Radiation Sciences for Oncology European Organization for Palliative Care European Organization for Research and Treatment of Cancer European Palliative Care Research Collaborative European School of Oncology European Society for Hyperthermic Oncology European Society for Intravenous Anaesthesia European Society for Medical Oncology European Society for Therapeutic Radiology and Oncology European Society of Anesthesiology European Society of Cancer Immunology and Immunotherapy European Society of Intensive Care Medicine European Society of Oncology Pharmacy European Society of Surgical Oncology Federación Panamericana e Ibérica de Sociedades de Medicina Crítica y Terapia Intensiva Fédération Nationale des Centres de Lutte Contre le Cancer Federation of Spanish Cancer Societies Fight Cancer Foundation Foundation for Anaesthesia Education and Research Foundation for European Education in Anaesthesiology Foundation of Geriatric Oncology Netherlands Freesia Group for Cancer Charities Spain French National Institute of Cancer French Society of Radiation Oncology French Society of Surgical Oncology German Cancer Aid German Cancer Research Center German Cancer Society German Society for Hematology and Oncology German Society of Radiation Oncology Ialian Association of Cancer Patients Intercultural Cancer Council Intercultural Cancer Council Caucus

International Agency for Research on Cancer International Anesthesia Research Society International Association for the Study of Pain International Cancer Biomarker Consortium International Cancer Microenvironment Society International Cancer Rehabilitation Association International Network for Cancer Treatment and Research International Organization for Cancer Prevention and International Society for Biological Therapy of Cancer International Society for Cell and Gene Therapy of Cancer International Society for Oncology and Biomarkers International Society of Cellular Oncology International Society of Intraoperative Radiation Therapy International Society of Oncology Pharmacy Practitioners International Union Against Cancer Ireland Cooperative Oncology Research Group Irish Association for Cancer Research Irish Association for Nurses in Oncology Irish Cancer Data Association Irish Cancer Society Irish Institute of Radiography and Radiation Therapy Irish Society of Medical Oncology Irish Society of Surgical Oncology Israel Cancer Association Italian Association for Cancer Research Italian Association for Radiation Oncology Italian Cancer Society Italian Foundation for Cancer Research Italian Institute for Cancer Rasearch and treatment Italian Institute of Medical Oncology Italian League Against Cancer Italian Society for Surgical Oncology Japan Clinical Cancer Research Organization Japan Society of Clinical Oncology Japan Society of Therapeutic Radiology and Oncology Japanese Cancer Association Japanese Foundation for Cancer Research Japanese Organization of Radiotherapy Quality Management Japanese Society of Anesthesiologists Japanese Society of Hyperthemic Oncology Japanese Society of Medical Oncology La Ligue Nationale contre le Cancer La Sociedad Española del Dolor La"Sociedad Española de Anestesiología, Reanimación y Terapéutica del Dolor l'Association Ensemble contre la douleur L'Association pour la Recherche sur le Cancer (ARC) Latin American and Caribbean Society of Medical Oncology Latin American Association for Palliative Care Latin American Cancer Research Coalition Macmillan Cancer Support Medical Oncology Group of Australia Mediterranean School of Oncology Multinational Association of Supportive Care in Cancer National Association of Professional Cancer Coaches National Cancer Institute National Cancer Registrars Association National Cancer Research Institute National Cancer Research Network National Coalition for Cancer Survivorship National Comprehensive Cancer Network National Foundation for Cancer Research National Health and Medical Research Council National Institute of Health and Excellence

Navy Anesthesia Society Nederlandse Vereniging voor Anesthesiologie New Zealand Society for Oncology New Zealand Society of Anaesthetists Nordic Cancer Union Norwegian Cancer Society Norwegian Group on Inherited Cancer Norwegian Society of Anaesthesiology Oncology Nutrition Dietetic Group Organisation of European Cancer Institutes Organization for Oncology and Translational Research Österreichische Gesellschaft für Internistische und Allgemeine Intensivmedizin Peripheral Regional Anesthesia Physician Assistants in Anesthesia Prevent Cancer Foundation Radiation Therapy Oncology Group Royal Australian & New Zealand College of Radiologists Royal College of Anaesthetists Schweizerische Gesellschaft für Intensivmedizin-Société Suisse de Médecine Intensive Scientific Association of Swiss Radiation Oncology Scottish Intercollegiate Guidelines Network Sino-American Network for Therapeutic Radiology and Oncology Sociedad Española de Enfermería Oncológica Sociedad Española de Medicina Intensiva, Crítica y Unidades Coronarias Società Italiana di Anestesia, Analgesia, Rianimazione e Terapia Intensiva Societé de Réanimation de Langue Francaise Société Française d'Anesthésie et de Réanimation Societe Francaise du cancer Société suisse d'anesthésiologie et de réanimation/Schweizerische Gesellschaft für Anästhesiologie und Reanimation Society for Ambulatory Anesthesia Society for Anesthesia and Resuscitation of Belgium Society for Education in Anesthesia Society for Education in Anesthesia Society for Integrative Oncology Society for the Advancement of Geriatric Anesthesia Society of Academic Anesthesiology Associations Society of Neurosurgical Anesthesia and Critical Care Society of Radiation Oncology Administrations Society of Surgical Oncology South African Oncology Consortium South African Society of Clinical and Radiation Oncology South African Society of Medical Oncology South East Asian Radiation Oncology Group (SEAROG) Southeast Anesthesiology Consultants Spanish Association Against Cancer Spanish Association for Cancer Research Spanish Association of Radiotherapy and Oncology Spanish Society of Chemotherapy Spanish Society of Medical Oncology: Spanish Society of Surgical Oncology Supportive and Rehabilitation Oncology Swedish Cancer Society Swedish Society for Anaesthesiology and Intensive Care Swedish Society of Oncology

Swedish Surgical Society Swiss Bridge Foundation Swiss Cancer League, Swiss League Against Cancer Swiss Cancer Research Foundation Swiss Federation Against Cancer (Oncosuisse) Swiss Group of Clinical Cancer Research Swiss Institute for Experimental Cancer Research Swiss Radiation Oncology Centers Swiss Society for Oncology Swiss Society of Medical Oncology Swiss Society of Surgery Taiwan Clinical Oncology Society The American Academy of Pain Medicine The American Board of Anesthesiology The American Academy of Anesthesiologist Assistants The American Chronic Pain Association The American College of Surgeons Oncology Group (ACOSOG The Anaesthesia Research Trust The Anesthesia Foundation The Association of Anaesthetists of Great Britain and Ireland The Association of Anesthesia Clinical Directors The Australian Organisation for Young People Living with Cancer The Australian Pain Society The Australian Patient Safety Foundation The Australian Society of Post Anaesthesia and Anaesthesia Nurses The Austrian Cancer league The Belgian Society of Intensive Care Medicine The British Medical Acupuncture Society The British Pain Society The Canadian Anesthesiologists' Society The Cancer Information and Support Society The European Cancer Patient Coalition The European Oncology Nursing Society The European Society of Digestive Oncology The European Society of Regional Anesthesia and Pain Therapy The Global Regional Anesthesia website The Intensive Care Society of Ireland The International Society for Anesthetic Pharmacology The International Spine Intervention Society The Japan Cancer Society The Japanese Association for Molecular Target Therapy of Cancer The National Board of Anesthesiology The Neuroanaesthesia Society of Great Britain and Ireland The New Zealand Association of Cancer Specialists The Royal College of Radiologists The Society of Anaesthetists of Hong Kong The South African Society of Anaesthesiologists The South Asian Association for Regional Cooperation The UK Society for Intravenous Society Trans Tasman Radiation Oncology Group World Anesthesia Society World Cancer Research Fund International World Federation Societies of Anesthesiologists World Federation of Surgical Oncology Societies

World Institute of Pain